



(ii) the APP<sub>695</sub> isoform of the APP gene having the specific mutations K670N, M671L and V717F; and

(iii) a transgenic non-human mammal whose genome comprises the gene of (ii).

Not only does Figure 2 fail to disclose the claimed invention, but nowhere in Hsaio is there any disclosure of a transgene encoding any APP isoform, let alone APP<sub>695</sub>, containing the specific mutations K670N, M671L and V717F. Hsaio's disclosure of transgenes encoding APP is limited to those disclosed in Table 2 (col. 2) and those reduced to practice (col. 22, ll. 31-49). Hsaio discloses only one transgene encoding an APP<sub>695</sub> with three mutations, i.e., the transgene encoding human APP<sub>695</sub> with the triple mutation V717I, V721IA, and M722V. *See, e.g.*, col. 22, ll. 43-44.

Because Hsaio fails to disclose each and every element of the claimed invention, applicants' respectfully request that the rejection be withdrawn and the claims passed to issuance.


### CONCLUSIONS

In view of the foregoing remarks, it is respectfully requested that the application be reconsidered and that all pending claims be allowed and the case passed to issue.

If there are any other issues remaining, which the Examiner believes could be resolved through either a Supplemental Response or an Examiner's Amendment, the Examiner is respectfully requested to contact the undersigned at the telephone number indicated below.

Respectfully submitted,

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